

REMARKS

Applicants have retained new counsel. Consequently, please **disregard** the Preliminary Amendment filed in connection with the above-captioned patent application on January 23, 2002.

The title has been amended as shown in Appendix A to more accurately reflect the subject matter now claimed. The specification has been amended as shown in Appendix A to correct an error in the naming of the claimed compound, which is discussed in detail below. The sequence listing has been replaced with the listing attached hereto as Appendix B. The attached sequence listing, which is identical to that submitted with the Preliminary Amendment filed January 23, 2002, has been resubmitted to avoid any confusion. In accordance with the requirements of 37 C.F.R. § 1.821(f), it is certified that the contents of the attached paper sequence listing and that of the computer readable copy submitted with this application are the same.

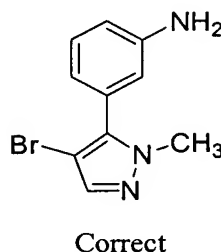
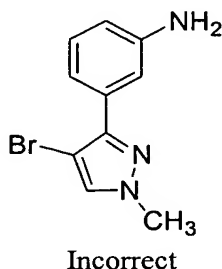
New claims 14-16 are pending in this application. Claims 14 and 15 are supported by the specification as filed. (*See* Experiment 15, page 73, lines 16-26). As discussed below, claim 16 is also supported by the application as filed.

It is well established that a claim may recite an inherent property of an invention described in an application, even if that application does not explicitly disclose the inherent property. *See Kennecott Corp. v. Kyocera International, Inc.*, 835 F.2d 1419, 1423 (Fed. Cir. 1987) ("The disclosure in a subsequent patent application of an inherent property of a product does not deprive that product of the benefit of an earlier filing date. Nor does the inclusion of a description of that property in later-filed claims change this reasonable result"). For example, the Court of Customs and Patent Appeals held on several occasions that the addition to an application of the chemical structure of a compound for which chemical properties had already been disclosed is not new matter. *See, e.g., In re Edwards*, 568 F.2d 1349 (CCPA 1978) (holding that a description of how to make a compound provided support for later filed claims that recited the compound itself); *In re Nathan*, 328 F.2d 1005, 1008 (CCPA 1964) (reversing a rejection of claims that recited the chemical orientation of a compound that was not explicitly described in the patent application, but which was an intrinsic property of the compound for which melting point, optical rotation, ultraviolet spectral analysis and chemical analysis data were provided).

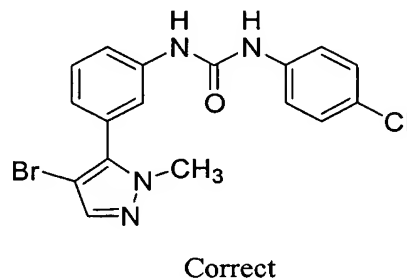
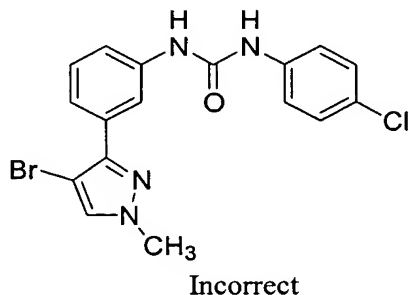
This application describes the synthesis of compounds that modulate serotonin receptors, and which may be useful in the treatment and prevention of a variety of diseases. (Page 3, lines 25-27). Spectroscopic and chromatographic properties of one of those

compounds are described in Experiment 15, which provides the basis for new claims 14 and 15. (Page 73, lines 16-26). In particular, claim 14 recites a serotonin receptor modulator having the ^1H NMR spectrum described in Experiment 15, and claim 15 recites a serotonin receptor modulator having the mass spectrum described in Experiment 15. New claim 16 recites a serotonin receptor modulator having the chemical structure of the compound described in Experiment 15. As discussed below, the correct chemical structure of that compound was recently discovered.

The compound described in Experiment 15 was made from a commercially available starting material obtained from Maybridge Chemical Company ("Maybridge"). (Page 55, lines 6-19). After the parent of this application (*i.e.*, application no. 09/292,072) was filed, Applicants discovered that the structure assigned the starting material by Maybridge was incorrect. In particular, it was discovered that the methyl group was attached to the other nitrogen atom of the pyrazole ring, as shown below:



Unfortunately, the incorrect structure was used to assign a structure to the compound of Experiment 15, which Applicants now realize was incorrect:



This realization is based, in part, on Applicants' preparation and testing of a compound that actually does have the structure shown above on the left. As expected, that compound does not possess the same spectroscopic, chemical or biological properties as the compound of the invention, *i.e.*, the compound described in Experiment 15.

It is a fundamental axiom of chemistry that the chemical structure of a compound is an intrinsic property of that compound. Indeed, a compound is defined by its chemical structure. Furthermore, the chemical structure of a compound dictates its physical, chemical and biological properties. For example, the chemical structure of a compound determines its ¹H NMR and mass spectra, its chromatographic behavior, and *in vitro* binding affinities. See, e.g., J. McMurry, Organic Chemistry, 411-413 (2nd ed., 1988).

Physical, chemical and biological properties of the compound described in Experiment 15 were measured by Applicants using standard techniques and equipment available to those of ordinary skill in the art, and were described in the application as filed. (See, e.g., page 66, lines 5-21). Those properties include the ¹H NMR and mass spectra of the compound (page 73, lines 22-25), its chromatographic behavior under well defined conditions (page 73, lines 20-21, 26; page 66, lines 13-16), and its biological activity as measured using various well defined *in vitro* assays (page 39, second entry in table; page 21, line 15 - page 24, line 7). Like its chemical structure, those properties are inherent properties of the compound described in Experiment 15.


The specification has been amended to correctly name the compound disclosed in Example 15, and new claim 16 recites the correct structure of the compound. Because the structure of the compound is an inherent property of it, and the name of the compound simply reflects that structure using standard nomenclature, Applicants respectfully submit that no new matter has been added by this preliminary amendment. See, e.g., *Kennecott Corp.*, 835 F.2d 1419; *In re Edwards*, 568 F.2d 1349; *In re Nathan*, 328 F.2d 1005.

[remainder of page intentionally left blank]

No fee is believed due for this submission. If one or more fees are due for this submission or to prevent the abandonment of the application, please charge such fee(s) to Pennie & Edmonds LLP Deposit Account No. 16-1150.

Respectfully submitted,

Date September 24, 2002

 45,479
Max Bachrach (Reg. No.)
PENNIE & EDMONDS LLP
1667 K Street, N.W., Suite 1000
Washington, DC 20006
(202) 496-4400

For: Laura A. Coruzzi (Reg. No. 30,742)
PENNIE & EDMONDS LLP
1155 Avenue of the Americas
New York, NY 10036-2711
(212) 790-9090

Attachments

APPENDIX A

Appendix A

Marked-up Version of Amendments to Application No. 10/055,555

In the Title:

Please amend the title as follows:

[NON-ENDOGENOUS, CONSTITUTIVELY ACTIVATED HUMAN] 2-METHYLPYRAZOLE BASED SEROTONIN [RECEPTORS AND SMALL MOLECULE] MODULATORS [THEREOF]

In the Specification:

Please amend the paragraph immediately following the title as follows:

[The benefit of U.S. Serial Number] This application is a continuation of application no. 09/292,072, filed April 14, 1999, which is a continuation-in-part of application no. 09/060,188, filed April 14, 1998 [(owned by Arena Pharmaceuticals, Inc.) and U.S. Provisional Number 60/090,783, filed June 26, 1998 (owned by Arena Pharmaceuticals), U.S. Provisional Number 60/112,909, filed December 18, 1998], and which claims priority to provisional application no. 60/123,000, filed March 5, 1999, provisional application no. 60/112,909, filed December 18, 1998, and provisional application no. [U.S. Provisional Number] 60/090,783, filed June 26, 1998.

Please amend the second chemical name provided in the table on page 39 (*i.e.*, the name provided in the third row of the table below its header) as follows: N-[3-(4-bromo-[1]2-methylpyrazol-3-yl)phenyl][(4-chlorophenyl)amino]-carboxamine

Please amend the chemical name provided on page 73, line 18, as follows: N-[3-(4-bromo-[1]2-methylpyrazol-3-yl)phenyl][(4-chlorophenyl)amino]-carboxamine

APPENDIX B

AREN0315.ST25.txt
SEQUENCE LISTING

RECEIVED
TECH CENTER 1600/2900
02 SEP 24 PM 12:12

<110> Behan, Dominic
Foster, Richard J.
Glen, Robert C.
Lawless, Michael S.
Liu, Qian
Smith, Julian R.
Liaw, Chen W.
Russo, Joseph F.
Thomsen, William J.
Chalmers, Derick

<120> Non-Endogenous, Constitutively Activated Human Serotonin Receptors and Small Molecule Modulators Thereof

<130> Aren-0315

<150> 09/292,072
<151> 1999-04-14

<150> 60/090,783
<151> 1998-06-26

<150> 60/112,909
<151> 1998-12-18

<150> 60/123,000
<151> 1999-03-05

<150> 09/060,188
<151> 1998-04-14

<150> 08/839,449
<151> 1997-04-14

<160> 33

<170> PatentIn version 3.1

<210> 1
<211> 27
<212> DNA
<213> Artificial Sequence

<220>
<223> Novel Sequence

<400> 1
gacctcgagg ttgcttaaga ctgaagc

27

<210> 2
<211> 27
<212> DNA
<213> Artificial Sequence

<220>
<223> Novel Sequence

<400> 2
atttctagac atatgtagct tgtaccg

27

<210> 3
<211> 50
<212> DNA

RECEIVED
SEP 26 2002
TECH CENTER 1600/2900

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 3

ctaggggcac catgcaggct atcaacaatg aaagaaaagc taagaaagtc

50

<210> 4

<211> 50

<212> DNA

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 4

caaggacttt cttagctttt ctttcattgt tgatagcctg catggtgccc

50

<210> 5

<211> 26

<212> DNA

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 5

gacctcgagt cttctacac ctcac

26

<210> 6

<211> 30

<212> DNA

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 6

tgctctagat tccagatagg tgaaaacttg

30

<210> 7

<211> 31

<212> DNA

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 7

caaagaaagt actgggcatc gtcttcttcc t

31

<210> 8

<211> 31

<212> DNA

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 8

ccgctcgagt actgcgccga caagctttga t

31

<210> 9
 <211> 38
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Novel Sequence

<400> 9
 cgatgccag cactttcgaa gcttttcttt cattgttg 38

<210> 10
 <211> 36
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Novel Sequence

<400> 10
 aaaagcttcg aaagtgctgg gcatcgtctt cttcct 36

<210> 11
 <211> 30
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Novel Sequence

<400> 11
 tgctctagat tccagatagg tgaaaacttg 30

<210> 12
 <211> 19
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Novel Sequence

<400> 12
 cgtgtctctc cttacttca 19

<210> 13
 <211> 36
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Novel Sequence

<400> 13
 tcggcgcagt actttgatag ttagaaagta ggtgat 36

<210> 14
 <211> 38
 <212> DNA
 <213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 14

ttctaactat caaagtactg cgccgacaag ctttgatg

38

<210> 15

<211> 43

<212> DNA

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 15

ttcagcagtc aaccactag tctatactct gttcaacaaa att

43

<210> 16

<211> 28

<212> DNA

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 16

atttctagac atatgtagct tgtaccgt

28

<210> 17

<211> 19

<212> DNA

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 17

atcacctact ttctaacta

19

<210> 18

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 18

ccataatcgt cagggaatg aaaaatgaca caa

33

<210> 19

<211> 33

<212> DNA

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 19

atttttcatt cccctgacga ttatggtgat tac

33

<210> 20
 <211> 33
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Novel Sequence

<400> 20
 tgatgaagaa agggcaccac atgatcagaa aca 33

<210> 21
 <211> 33
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Novel Sequence

<400> 21
 gatcatgtgg tgccctttct tcatcacaaa cat 33

<210> 22
 <211> 24
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Novel Sequence

<400> 22
 gagacatatt atctgccacg gagg 24

<210> 23
 <211> 24
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Novel Sequence

<400> 23
 ttggcataga aaccggaccc aagg 24

<210> 24
 <211> 1416
 <212> DNA
 <213> Artificial Sequence

<220>
 <223> Novel Sequence

<400> 24
 atggatattc tttgtgaaga aaatacttct ttgagctcaa ctacgaactc cctaattgcaa 60
 ttaaatgatg acaacaggct ctacagtaat gactttaact ccggagaagc taacacttct 120
 gatgcattta actggacagt cgactctgaa aatcgaacca acctttcctg tgaaggggtgc 180
 ctctcaccgt cgtgtctctc cttacttcat ctccaggaaa aaaactggtc tgctttactg 240
 acagccgtag tgattattct aactattgct ggaaacatac tcgtcatcat ggcagtggtcc 300

AREN0315.ST25.txt

ctagagaaaa agctgcagaa tgccaccaac tatttcctga tgtcacttgc catagctgat 360
atgctgctgg gtttccttgt catgcccgtg tccatgttaa ccatcctgta tgggtaccgg 420
tggcctctgc cgagcaagct ttgtgcagtc tggatttacc tggacgtgct cttctccacg 480
gcctccatca tgcacctctg cgccatctcg ctggaccgct acgtcgccat ccagaatccc 540
atccaccaca gccgcttcaa ctccagaact aaggcatttc tgaaaatcat tgctgtttgg 600
accatatcag taggtatata catgccata ccagtctttg ggctacagga cgattcgaag 660
gtctttaagg aggggagttg cttactcgcc gatgataact ttgtcctgat cggctctttt 720
gtgtcatttt tcattccctt aaccatcatg gtgatcacct actttctaac tatcaagtca 780
ctccagaaag aagctacttt gtgtgtaagt gatcttggca cacgggcaa attagcttct 840
ttcagcttcc tccctcagag ttctttgtct tcagaaaagc tcttcacgag gtcgatccat 900
agggagccag ggtcctacac aggcaggagg actatgcagt ccatcagcaa tgagcaaaag 960
gcatgcaagg tgctgggcat cgtcttcttc ctgtttgtgg tgatgtggtg ccctttcttc 1020
atcacaacaa tcatggccgt catctgcaaa gagtcttgca atgaggatgt cattggggcc 1080
ctgctcaatg tgtttgtttg gatcggttat ctctcttcag cagtcaaccc actagtctac 1140
acactgttca acaagaccta taggtcagcc ttttcacggt atattcagtg tcagtacaag 1200
gaaaacaaaa aaccattgca gttaatttta gtgaacacaa taccggcttt ggcctacaag 1260
tctagccaac ttcaaattgg acaaaaaaag aattcaaagc aagatgcaa gacaacagat 1320
aatgactgct caatggttgc tctaggaaag cagtattctg aagaggcttc taaagacaat 1380
agcgacggag tgaatgaaaa ggtgagctgt gtgtga 1416

<210> 25
<211> 470
<212> PRT
<213> Artificial Sequence

<220>
<223> Novel Sequence

<400> 25

Met Asp Ile Leu Cys Glu Glu Asn Thr Ser Leu Ser Ser Thr Thr Asn
1 5 10 15

Ser Leu Met Gln Leu Asn Asp Asp Asn Arg Leu Tyr Ser Asn Asp Phe
20 25 30

Asn Ser Gly Glu Ala Asn Thr Ser Asp Ala Phe Asn Trp Thr Val Asp
35 40 45

Ser Glu Asn Arg Thr Asn Leu Ser Cys Glu Gly Cys Leu Ser Pro Ser
50 55 60

Cys Ser Leu Leu His Leu Gln Glu Lys Asn Trp Ser Ala Leu Leu Thr
65 70 75 80

AREN0315.ST25.txt

Ala Val Val Ile Ile Leu Thr Ile Ala Gly Asn Ile Leu Val Ile Met
85 90 95

Ala Val Ser Leu Glu Lys Lys Leu Gln Asn Ala Thr Asn Tyr Phe Leu
100 105 110

Met Ser Leu Ala Ile Ala Asp Met Leu Leu Gly Phe Leu Val Met Pro
115 120 125

Val Ser Met Leu Thr Ile Leu Tyr Gly Tyr Arg Trp Pro Leu Pro Ser
130 135 140

Lys Leu Cys Ala Val Trp Ile Tyr Leu Asp Val Leu Phe Ser Thr Ala
145 150 155 160

Ser Ile Met His Leu Cys Ala Ile Ser Leu Asp Arg Tyr Val Ala Ile
165 170 175

Gln Asn Pro Ile His His Ser Arg Phe Asn Ser Arg Thr Lys Ala Phe
180 185 190

Leu Lys Ile Ile Ala Val Trp Thr Ile Ser Val Gly Ile Ser Met Pro
195 200 205

Ile Pro Val Phe Gly Leu Gln Asp Asp Ser Lys Val Phe Lys Glu Gly
210 215 220

Ser Cys Leu Leu Ala Asp Asp Asn Phe Val Leu Ile Gly Ser Phe Val
225 230 235 240

Ser Phe Phe Ile Pro Leu Thr Ile Met Val Ile Thr Tyr Phe Leu Thr
245 250 255

Ile Lys Ser Leu Gln Lys Glu Ala Thr Leu Cys Val Ser Asp Leu Gly
260 265 270

Thr Arg Ala Lys Leu Ala Ser Phe Ser Phe Leu Pro Gln Ser Ser Leu
275 280 285

Ser Ser Glu Lys Leu Phe Gln Arg Ser Ile His Arg Glu Pro Gly Ser
290 295 300

Tyr Thr Gly Arg Arg Thr Met Gln Ser Ile Ser Asn Glu Gln Lys Ala
305 310 315 320

Cys Lys Val Leu Gly Ile Val Phe Phe Leu Phe Val Val Met Trp Cys
325 330 335

Pro Phe Phe Ile Thr Asn Ile Met Ala Val Ile Cys Lys Glu Ser Cys

340

345

350

Asn Glu Asp Val Ile Gly Ala Leu Leu Asn Val Phe Val Trp Ile Gly
 355 360 365

Tyr Leu Ser Ser Ala Val Asn Pro Leu Val Tyr Thr Leu Phe Asn Lys
 370 375 380

Thr Tyr Arg Ser Ala Phe Ser Arg Tyr Ile Gln Cys Gln Tyr Lys Glu
 385 390 395 400

Asn Lys Lys Pro Leu Gln Leu Ile Leu Val Asn Thr Ile Pro Ala Leu
 405 410 415

Ala Tyr Lys Ser Ser Gln Leu Gln Met Gly Gln Lys Lys Asn Ser Lys
 420 425 430

Gln Asp Ala Lys Thr Thr Asp Asn Asp Cys Ser Met Val Ala Leu Gly
 435 440 445

Lys Gln Tyr Ser Glu Glu Ala Ser Lys Asp Asn Ser Asp Gly Val Asn
 450 455 460

Glu Lys Val Ser Cys Val
 465 470

<210> 26

<211> 1377

<212> DNA

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 26

atgggtgaacc tgaggaatgc ggtgcattca ttccttgtgc acctaattgg cctattgggt 60
 tggcaatgtg atatttctgt gagcccagta gcagctatag taactgacat tttcaatacc 120
 tccgatggtg gacgcttcaa attcccagac ggggtacaaa actggccagc actttcaatc 180
 gtcatacataa taatcatgac aataggtggc aacatccttg tgatcatggc agtaagcatg 240
 gaaaagaaac tgcacaatgc caccaattac ttcttaatgt ccctagccat tgctgatatg 300
 ctagtgggac tacttgtcat gcccctgtct ctctggcaa tcctttatga ttatgtctgg 360
 ccactaccta gatatttgtg ccccgctctg atttcttttag atgttttatt ttcaacagcg 420
 tccatcatgc acctctgcgc tatatcgctg gatcggtatg tagcaatacg taatcctatt 480
 gagcatagcc gtttcaattc gcggactaag gccatcatga agattgctat tgtttgggca 540
 atttctatag gtgtatcagt tcctatccct gtgattggac tgagggacga agaaaagggtg 600
 ttcgtgaaca acacgacgtg cgtgctcaac gacccaaatt tcgttcttat tgggtccttc 660
 gtagctttct tcataccgct gacgattatg gtgattacgt attgcctgac catctacgtt 720

AREN0315.ST25.txt

ctgcgccgac aagctttgat gttactgcac ggccacaccg aggaaccgcc tggactaagt 780
ctggatttcc tgaagtgctg caagaggaat acggccgagg aagagaactc tgcaaaccct 840
aaccaagacc agaacgcacg ccgaagaaag aagaaggaga gacgtcctag gggcaccatg 900
caggctatca acaatgaaag aaaagcttcg aaagtccttg ggattgtttt ctttgtgttt 960
ctgatcatgt ggtgccccatt ttccattacc aatattctgt ctgttctttg tgagaagtcc 1020
tgtaaccaaa agctcatgga aaagcttctg aatgtgtttg ttggattgg ctatgtttgt 1080
tcaggaatca atcctctggt gtatctctgt ttcaacaaaa ttaccgaag ggcattctcc 1140
aactatttgc gttgcaatta taaggtagag aaaaagcctc ctgtcaggca gattccaaga 1200
gttgccgccca ctgctttgtc tgggaggagg cttaatgtta acatttatcg gcataccaat 1260
gaaccggtga tcgagaaagc cagtgacaat gagcccggta tagagatgca agttgagaat 1320
ttagagttac cagtaaatac ctccagtgtg gttagcgaaa ggattagcag tgtgtga 1377

<210> 27
<211> 458
<212> PRT
<213> Artificial Sequence

<220>
<223> Novel Sequence

<400> 27

Met Val Asn Leu Arg Asn Ala Val His Ser Phe Leu Val His Leu Ile
1 5 10 15

Gly Leu Leu Val Trp Gln Cys Asp Ile Ser Val Ser Pro Val Ala Ala
20 25 30

Ile Val Thr Asp Ile Phe Asn Thr Ser Asp Gly Gly Arg Phe Lys Phe
35 40 45

Pro Asp Gly Val Gln Asn Trp Pro Ala Leu Ser Ile Val Ile Ile Ile
50 55 60

Ile Met Thr Ile Gly Gly Asn Ile Leu Val Ile Met Ala Val Ser Met
65 70 75 80

Glu Lys Lys Leu His Asn Ala Thr Asn Tyr Phe Leu Met Ser Leu Ala
85 90 95

Ile Ala Asp Met Leu Val Gly Leu Leu Val Met Pro Leu Ser Leu Leu
100 105 110

Ala Ile Leu Tyr Asp Tyr Val Trp Pro Leu Pro Arg Tyr Leu Cys Pro
115 120 125

Val Trp Ile Ser Leu Asp Val Leu Phe Ser Thr Ala Ser Ile Met His

130

135

Leu Cys Ala Ile Ser Leu Asp Arg Tyr Val Ala Ile Arg Asn Pro Ile
145 150 155 160

Glu His Ser Arg Phe Asn Ser Arg Thr Lys Ala Ile Met Lys Ile Ala
165 170 175

Ile Val Trp Ala Ile Ser Ile Gly Val Ser Val Pro Ile Pro Val Ile
180 185 190

Gly Leu Arg Asp Glu Glu Lys Val Phe Val Asn Asn Thr Thr Cys Val
195 200 205

Leu Asn Asp Pro Asn Phe Val Leu Ile Gly Ser Phe Val Ala Phe Phe
210 215 220

Ile Pro Leu Thr Ile Met Val Ile Thr Tyr Cys Leu Thr Ile Tyr Val
225 230 235 240

Leu Arg Arg Gln Ala Leu Met Leu Leu His Gly His Thr Glu Glu Pro
245 250 255

Pro Gly Leu Ser Leu Asp Phe Leu Lys Cys Cys Lys Arg Asn Thr Ala
260 265 270

Glu Glu Glu Asn Ser Ala Asn Pro Asn Gln Asp Gln Asn Ala Arg Arg
275 280 285

Arg Lys Lys Lys Glu Arg Arg Pro Arg Gly Thr Met Gln Ala Ile Asn
290 295 300

Asn Glu Arg Lys Ala Ser Lys Val Leu Gly Ile Val Phe Phe Val Phe
305 310 315 320

Leu Ile Met Trp Cys Pro Phe Phe Ile Thr Asn Ile Leu Ser Val Leu
325 330 335

Cys Glu Lys Ser Cys Asn Gln Lys Leu Met Glu Lys Leu Leu Asn Val
340 345 350

Phe Val Trp Ile Gly Tyr Val Cys Ser Gly Ile Asn Pro Leu Val Tyr
355 360 365

Thr Leu Phe Asn Lys Ile Tyr Arg Arg Ala Phe Ser Asn Tyr Leu Arg
370 375 380

Cys Asn Tyr Lys Val Glu Lys Lys Pro Pro Val Arg Gln Ile Pro Arg
385 390 395 400

AREN0315.ST25.txt

Val Ala Ala Thr Ala Leu Ser Gly Arg Glu Leu Asn Val Asn Ile Tyr
405 410 415

Arg His Thr Asn Glu Pro Val Ile Glu Lys Ala Ser Asp Asn Glu Pro
420 425 430

Gly Ile Glu Met Gln Val Glu Asn Leu Glu Leu Pro Val Asn Pro Ser
435 440 445

Ser Val Val Ser Glu Arg Ile Ser Ser Val
450 455

<210> 28
<211> 1377
<212> DNA
<213> Artificial Sequence

<220>
<223> Novel Sequence

<400> 28
atggtgaacc tgaggaatgc ggtgcattca ttccttgtgc acctaattgg cctattggtt 60
tggaatgtg atatttctgt gagccagta gcagctatag taactgacat tttcaatacc 120
tccgatggtg gacgcttcaa attcccagac ggggtacaaa actggccagc actttcaatc 180
gtcatcataa taatcatgac aataggtggc aacatccttg tgatcatggc agtaagcatg 240
gaaaagaaac tgcacaatgc caccaattac ttcttaatgt ccctagccat tgctgatatg 300
ctagtgggac tacttgtcat gccctgtct ctctggcaa tcctttatga ttatgtctgg 360
ccactaccta gatatttgtg cccgtcttg atttcttag atgttttatt ttcaacagcg 420
tccatcatgc acctctgcgc tatatcgctg gatcggtatg tagcaatacg taatcctatt 480
gagcatagcc gtttcaattc gcggactaag gccatcatga agattgctat tgtttgggca 540
atttctatag gtgtatcagt tcctatccct gtgattggac tgagggacga agaaaagggtg 600
ttcgtgaaca acacgacgtg cgtgctcaac gacccaaatt tcgttcttat tgggtccttc 660
gtagctttct tcataccgct gacgattatg gtgattacgt attgcctgac catctacgtt... 720
ctgcgccgac aagctttgat gttactgcac ggccacaccg aggaaccgcc tggactaagt 780
ctggatttcc tgaagtgtg caagaggaat acggccgagg aagagaactc tgcaaaccct 840
aaccaagacc agaacgcacg ccgaagaaag aagaaggaga gacgtcctag gggcaccatg 900
caggctatca acaatgaaag aaaagctaag aaagtccttg ggattgtttt ctttgtgttt 960
ctgatcatgt ggtgcccatt ttctattacc aatattctgt ctgttctttg tgagaagtcc 1020
tgtaacaaa agctcatgga aaagcttctg aatgtgtttg tttggattgg ctatgtttgt 1080
tcaggaatca atcctctggt gtatactctg ttcaacaaaa tttaccgaag ggcattctcc 1140
aactatttgc gttgcaatta taaggtagag aaaaagcctc ctgtcaggca gattccaaga 1200
gttgccgcca ctgctttgtc tgggaggagg cttaatgtta acatttatcg gcataccaat 1260

AREN0315.ST25.txt

gaaccggtga tcgagaaagc cagtgacaat gagcccgta tagagatgca agttgagaat 1320
ttagagttac cagtaaattcc ctccagtgtg gtttagcgaaa ggattagcag tgtgtga 1377

<210> 29
<211> 458
<212> PRT
<213> Artificial Sequence

<220>
<223> Novel Sequence

<400> 29

Met Val Asn Leu Arg Asn Ala Val His Ser Phe Leu Val His Leu Ile
1 5 10 15

Gly Leu Leu Val Trp Gln Cys Asp Ile Ser Val Ser Pro Val Ala Ala
20 25 30

Ile Val Thr Asp Ile Phe Asn Thr Ser Asp Gly Gly Arg Phe Lys Phe
35 40 45

Pro Asp Gly Val Gln Asn Trp Pro Ala Leu Ser Ile Val Ile Ile Ile
50 55 60

Ile Met Thr Ile Gly Gly Asn Ile Leu Val Ile Met Ala Val Ser Met
65 70 75 80

Glu Lys Lys Leu His Asn Ala Thr Asn Tyr Phe Leu Met Ser Leu Ala
85 90 95

Ile Ala Asp Met Leu Val Gly Leu Leu Val Met Pro Leu Ser Leu Leu
100 105 110

Ala Ile Leu Tyr Asp Tyr Val Trp Pro Leu Pro Arg Tyr Leu Cys Pro
115 120 125

Val Trp Ile Ser Leu Asp Val Leu Phe Ser Thr Ala Ser Ile Met His
130 135 140

Leu Cys Ala Ile Ser Leu Asp Arg Tyr Val Ala Ile Arg Asn Pro Ile
145 150 155 160

Glu His Ser Arg Phe Asn Ser Arg Thr Lys Ala Ile Met Lys Ile Ala
165 170 175

Ile Val Trp Ala Ile Ser Ile Gly Val Ser Val Pro Ile Pro Val Ile
180 185 190

Gly Leu Arg Asp Glu Glu Lys Val Phe Val Asn Asn Thr Thr Cys Val
195 200 205

AREN0315.ST25.txt

Leu Asn Asp Pro Asn Phe Val Leu Ile Gly Ser Phe Val Ala Phe Phe
210 215 220

Ile Pro Leu Thr Ile Met Val Ile Thr Tyr Cys Leu Thr Ile Tyr Val
225 230 235 240

Leu Arg Arg Gln Ala Leu Met Leu Leu His Gly His Thr Glu Glu Pro
245 250 255

Pro Gly Leu Ser Leu Asp Phe Leu Lys Cys Cys Lys Arg Asn Thr Ala
260 265 270

Glu Glu Glu Asn Ser Ala Asn Pro Asn Gln Asp Gln Asn Ala Arg Arg
275 280 285

Arg Lys Lys Lys Glu Arg Arg Pro Arg Gly Thr Met Gln Ala Ile Asn
290 295 300

Asn Glu Arg Lys Ala Lys Lys Val Leu Gly Ile Val Phe Phe Val Phe
305 310 315 320

Leu Ile Met Trp Cys Pro Phe Phe Ile Thr Asn Ile Leu Ser Val Leu
325 330 335

Cys Glu Lys Ser Cys Asn Gln Lys Leu Met Glu Lys Leu Leu Asn Val
340 345 350

Phe Val Trp Ile Gly Tyr Val Cys Ser Gly Ile Asn Pro Leu Val Tyr
355 360 365

Thr Leu Phe Asn Lys Ile Tyr Arg Arg Ala Phe Ser Asn Tyr Leu Arg
370 375 380

Cys Asn Tyr Lys Val Glu Lys Lys Pro Pro Val Arg Gln Ile Pro Arg
385 390 395 400

Val Ala Ala Thr Ala Leu Ser Gly Arg Glu Leu Asn Val Asn Ile Tyr
405 410 415

Arg His Thr Asn Glu Pro Val Ile Glu Lys Ala Ser Asp Asn Glu Pro
420 425 430

Gly Ile Glu Met Gln Val Glu Asn Leu Glu Leu Pro Val Asn Pro Ser
435 440 445

Ser Val Val Ser Glu Arg Ile Ser Ser Val
450 455

<210> 30
<211> 1437
<212> DNA

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 30

```

atggatattc tttgtgaaga aaatacttct ttgagctcaa ctacgaactc cctaatagcaa    60
ttaaatagatg acaacaggct ctacagtaat gactttaact ccggagaagc taacacttct    120
gatgcattta actggacagt cgactctgaa aatcgaacca acctttcctg tgaaggggtgc    180
ctctcaccgt cgtgtctctc cttacttcat ctccaggaaa aaaactggtc tgctttactg    240
acagccgtag tgattattct aactattgct ggaacatac tcgtcatcat ggcagtgtcc    300
ctagagaaaa agctgcagaa tgccaccaac tatttcctga tgtcacttgc catagctgat    360
atgctgctgg gtttccttgt catgcccggtg tccatgttaa ccatcctgta tgggtaccgg    420
tggcctctgc cgagcaagct ttgtgcagtc tggatttacc tggacgtgct cttctccacg    480
gcctccatca tgcacctctg cgccatctcg ctggaccgct acgtcgccat ccagaatccc    540
atccaccaca gccgcttcaa ctccagaact aaggcatttc tgaaaatcat tgctgtttgg    600
accatatcag taggtatata catgccataa ccagtctttg ggctacagga cgattcgaag    660
gtctttaagg aggggagttg cttactcgcc gatgataact ttgtcctgat cggctctttt    720
gtgtcatttt tcattccctt aaccatcatg gtgatcacct actttctaac tatcaaggtt    780
ctgcgccgac aagctttgat gttactgcac ggccacaccg aggaaccgcc tggactaagt    840
ctggatttcc tgaagtgtg caagaggaat acggccgagg aagagaactc tgcaaaccct    900
aaccaagacc agaacgcacg ccgaagaaag aagaaggaga gacgtcctag gggcaccatg    960
caggctatca acaatgaaag aaaagcttcg aaggtagtgg gcatcgtctt cttcctgttt   1020
gtgggtgatgt ggtgcccttt cttcatcaca aacatcatgg ccgtcatctg caaagagtcc   1080
tgcaatgagg atgtcattgg ggccctgctc aatgtgtttg tttggatcgg ttatctctct   1140
tcagcagtca acccactagt ctatactctg ttcaacaaaa tttaccgaag ggcattctcc   1200
aactatttgc gttgcaatta taaggtagag aaaaagcctc ctgtcaggca gattccaaga   1260
gttgccgccca ctgctttgtc tgggagggag cttaatgtta acatttatcg gcataccaat   1320
gaaccggtga tcgagaaagc cagtgacaat gagcccggtg tagagatgca agttgagaat   1380
ttagagttac cagtaaatcc ctccagtgtg gttagcgaaa ggattagcag tgtgtga     1437

```

<210> 31

<211> 478

<212> PRT

<213> Artificial Sequence

<220>

<223> Novel Sequence

<400> 31

```

Met Asp Ile Leu Cys Glu Glu Asn Thr Ser Leu Ser Ser Thr Thr Asn
1           5           10           15

```

Ser Leu Met Gln Leu Asn Asp Asp Asn Arg Leu Tyr Ser Asn Asp Phe
20 25 30

Asn Ser Gly Glu Ala Asn Thr Ser Asp Ala Phe Asn Trp Thr Val Asp
35 40 45

Ser Glu Asn Arg Thr Asn Leu Ser Cys Glu Gly Cys Leu Ser Pro Ser
50 55 60

Cys Leu Ser Leu Leu His Leu Gln Glu Lys Asn Trp Ser Ala Leu Leu
65 70 75 80

Thr Ala Val Val Ile Ile Leu Thr Ile Ala Gly Asn Ile Leu Val Ile
85 90 95

Met Ala Val Ser Leu Glu Lys Lys Leu Gln Asn Ala Thr Asn Tyr Phe
100 105 110

Leu Met Ser Leu Ala Ile Ala Asp Met Leu Leu Gly Phe Leu Val Met
115 120 125

Pro Val Ser Met Leu Thr Ile Leu Tyr Gly Tyr Arg Trp Pro Leu Pro
130 135 140

Ser Lys Leu Cys Ala Val Trp Ile Tyr Leu Asp Val Leu Phe Ser Thr
145 150 155 160

Ala Ser Ile Met His Leu Cys Ala Ile Ser Leu Asp Arg Tyr Val Ala
165 170 175

Ile Gln Asn Pro Ile His His Ser Arg Phe Asn Ser Arg Thr Lys Ala
180 185 190

Phe Leu Lys Ile Ile Ala Val Trp Thr Ile Ser Val Gly Ile Ser Met
195 200 205

Pro Ile Pro Val Phe Gly Leu Gln Asp Asp Ser Lys Val Phe Lys Glu
210 215 220

Gly Ser Cys Leu Leu Ala Asp Asp Asn Phe Val Leu Ile Gly Ser Phe
225 230 235 240

Val Ser Phe Phe Ile Pro Leu Thr Ile Met Val Ile Thr Tyr Phe Leu
245 250 255

Thr Ile Lys Val Leu Arg Arg Gln Ala Leu Met Leu Leu His Gly His
260 265 270

Thr Glu Glu Pro Pro Gly Leu Ser Leu Asp Phe Leu Lys Cys Cys Lys

275

280

Arg Asn Thr Ala Glu Glu Glu Asn Ser Ala Asn Pro Asn Gln Asp Gln
290 295 300

Asn Ala Arg Arg Arg Lys Lys Lys Glu Arg Arg Pro Arg Gly Thr Met
305 310 315 320

Gln Ala Ile Asn Asn Glu Arg Lys Ala Ser Lys Val Leu Gly Ile Val
325 330 335

Phe Phe Leu Phe Val Val Met Trp Cys Pro Phe Phe Ile Thr Asn Ile
340 345 350

Met Ala Val Ile Cys Lys Glu Ser Cys Asn Glu Asp Val Ile Gly Ala
355 360 365

Leu Leu Asn Val Phe Val Trp Ile Gly Tyr Leu Ser Ser Ala Val Asn
370 375 380

Pro Leu Val Tyr Thr Leu Phe Asn Lys Ile Tyr Arg Arg Ala Phe Ser
385 390 395 400

Asn Tyr Leu Arg Cys Asn Tyr Lys Val Glu Lys Lys Pro Pro Val Arg
405 410 415

Gln Ile Pro Arg Val Ala Ala Thr Ala Leu Ser Gly Arg Glu Leu Asn
420 425 430

Val Asn Ile Tyr Arg His Thr Asn Glu Pro Val Ile Glu Lys Ala Ser
435 440 445

Asp Asn Glu Pro Gly Ile Glu Met Gln Val Glu Asn Leu Glu Leu Pro
450 455 460

Val Asn Pro Ser Ser Val Val Ser Glu Arg Ile Ser Ser Val
465 470 475

<210> 32
<211> 1437
<212> DNA
<213> Artificial Sequence

<220>
<223> Novel Sequence

<400> 32
atggatattc tttgtgaaga aaatacttct ttgagctcaa ctacgaactc cctaattgcaa 60
ttaaatgatg acaacaggct ctacagtaat gactttaact ccggagaagc taacacttct 120
gatgcattta actggacagt cgactctgaa aatcgaacca acctttcctg tgaaggggtgc 180
ctctcaccgt cgtgtctctc cttacttcat ctccaggaaa aaaactggtc tgctttactg 240

AREN0315.ST25.txt

acagccgtag tgattattct aactattgct ggaaacatac tcgtcatcat ggcagtgtcc 300
ctagagaaaa agctgcagaa tgccaccaac tatttcctga tgtcacttgc catagctgat 360
atgctgctgg gtttccttgt catgcccgty tccatgttaa ccatcctgta tgggtacegg. 420
tggcctctgc cgagcaagct ttgtgcagtc tggatttacc tggacgtgct cttctccacg 480
gcctccatca tgcacctctg cgccatctcg ctggaccgct acgtcgccat ccagaatccc 540
atccaccaca gccgcttcaa ctccagaact aaggcatttc tgaaaatcat tgctgtttgg 600
accatatcag taggtatatc catgccaaata ccagtccttg ggctacagga cgattcgaag 660
gtctttaagg aggggagttg cttactcgcc gatgataact ttgtcctgat cggctctttt 720
gtgtcatttt tcattcccct gacgattatg gtgattacgt attgcctgac catctacgtt 780
ctgcgccgac aagctttgat gttactgcac ggccacaccg aggaaccgcc tggactaagt 840
ctggatttcc tgaagtgtg caagaggaat acggccgagg aagagaactc tgcaaaccct 900
aaccaagacc agaacgcacg ccgaagaaag aagaaggaga gacgtcctag gggcaccatg 960
caggctatca acaatgaaag aaaagctaag aaagtccttg ggattgtttt ctttgtgttt 1020
ctgatcatgt ggtgcccttt cttcatcaca aacatcatgg ccgtcatctg caaagagtcc 1080
tgcaatgagg atgtcattgg ggccctgctc aatgtgtttg tttggatcgg ttatctctct 1140
tcagcagtca acccactagt ctatactctg ttcaacaaaa tttaccgaag ggcattctcc 1200
aactatttgc gttgcaatta taaggtagag aaaaagcctc ctgtcaggca gattccaaga 1260
gttgccgcca ctgctttgtc tgggagggag cttaatgtta acatttatcg gcataccaat 1320
gaaccggtga tcgagaaagc cagtgcacat gagcccggtg tagagatgca agttgagaat 1380
ttagagttac cagtaaatac ctccagtgtg gtagcgaaa ggattagcag tgtgtga 1437

<210> 33
<211> 478
<212> PRT
<213> Artificial Sequence

<220>
<223> Novel Sequence

<400> 33

Met Asp Ile Leu Cys Glu Glu Asn Thr Ser Leu Ser Ser Thr Thr Asn
1 5 10 15

Ser Leu Met Gln Leu Asn Asp Asp Asn Arg Leu Tyr Ser Asn Asp Phe
20 25 30

Asn Ser Gly Glu Ala Asn Thr Ser Asp Ala Phe Asn Trp Thr Val Asp
35 40 45

Ser Glu Asn Arg Thr Asn Leu Ser Cys Glu Gly Cys Leu Ser Pro Ser
50 55 60

AREN0315.ST25.txt

Cys Leu Ser Leu Leu His Leu Gln Glu Lys Asn Trp Ser Ala Leu Leu
65 70 75 80

Thr Ala Val Val Ile Ile Leu Thr Ile Ala Gly Asn Ile Leu Val Ile
85 90 95

Met Ala Val Ser Leu Glu Lys Lys Leu Gln Asn Ala Thr Asn Tyr Phe
100 105 110

Leu Met Ser Leu Ala Ile Ala Asp Met Leu Leu Gly Phe Leu Val Met
115 120 125

Pro Val Ser Met Leu Thr Ile Leu Tyr Gly Tyr Arg Trp Pro Leu Pro
130 135 140

Ser Lys Leu Cys Ala Val Trp Ile Tyr Leu Asp Val Leu Phe Ser Thr
145 150 155 160

Ala Ser Ile Met His Leu Cys Ala Ile Ser Leu Asp Arg Tyr Val Ala
165 170 175

Ile Gln Asn Pro Ile His His Ser Arg Phe Asn Ser Arg Thr Lys Ala
180 185 190

Phe Leu Lys Ile Ile Ala Val Trp Thr Ile Ser Val Gly Ile Ser Met
195 200 205

Pro Ile Pro Val Phe Gly Leu Gln Asp Asp Ser Lys Val Phe Lys Glu
210 215 220

Gly Ser Cys Leu Leu Ala Asp Asp Asn Phe Val Leu Ile Gly Ser Phe
225 230 235 240

Val Ser Phe Phe Ile Pro Leu Thr Ile Met Val Ile Thr Tyr Cys Leu
245 250 255

Thr Ile Tyr Val Leu Arg Arg Gln Ala Leu Met Leu Leu His Gly His
260 265 270

Thr Glu Glu Pro Pro Gly Leu Ser Leu Asp Phe Leu Lys Cys Cys Lys
275 280 285

Arg Asn Thr Ala Glu Glu Glu Asn Ser Ala Asn Pro Asn Gln Asp Gln
290 295 300

Asn Ala Arg Arg Arg Lys Lys Lys Glu Arg Arg Pro Arg Gly Thr Met
305 310 315 320

Gln Ala Ile Asn Asn Glu Arg Lys Ala Lys Lys Val Leu Gly Ile Val
325 330 335

Phe Phe Val Phe Leu Ile Met Trp Cys Pro Phe Phe Ile Thr Asn Ile
340 345 350

Met Ala Val Ile Cys Lys Glu Ser Cys Asn Glu Asp Val Ile Gly Ala
355 360 365

Leu Leu Asn Val Phe Val Trp Ile Gly Tyr Leu Ser Ser Ala Val Asn
370 375 380

Pro Leu Val Tyr Thr Leu Phe Asn Lys Ile Tyr Arg Arg Ala Phe Ser
385 390 395 400

Asn Tyr Leu Arg Cys Asn Tyr Lys Val Glu Lys Lys Pro Pro Val Arg
405 410 415

Gln Ile Pro Arg Val Ala Ala Thr Ala Leu Ser Gly Arg Glu Leu Asn
420 425 430

Val Asn Ile Tyr Arg His Thr Asn Glu Pro Val Ile Glu Lys Ala Ser
435 440 445

Asp Asn Glu Pro Gly Ile Glu Met Gln Val Glu Asn Leu Glu Leu Pro
450 455 460

Val Asn Pro Ser Ser Val Val Ser Glu Arg Ile Ser Ser Val
465 470 475